## PATENT COOPERATION TREATY

To: ANNETTE S. PARENT TOWNSEND AND TOWNSEND AND CREW LLP TWO EMBARCADERO CENTER, 8TH FLOOR SAN FRANCISCO, CA 94111	PCT  NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT AND THE WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY, OR THE DECLARATION			
02307E-143900PC	(PCT Rule 44.1)  Date of mailing (day/month/year) 18 JUL 2005			
Applicant's or agent's file reference 2307E-1439PC	FOR FURTHER ACTION See paragraphs 1 and 4 below			
International application No. PCT/US05/07297	International filing date (day/month/year) 07 March 2005 (07.03.2005)			
Applicant THE REGENTS OF HTE UNIVERSITY OF CALIFORNIA				
The applicant is hereby notified that the international s     have been established and are transmitted herewith.	earch report and the written opinion of the International Searching Authority			
Filing of amendments and statement under Article The applicant is entitled, if he so wishes, to amend the	19:			
	s is normally two months from the date of transmittal of the international			
Where? Directly to the International Bureau of WI 1211 Geneva 20, Switzerland, Facsimile I	IPO, 34 chemin des Colombettes No.: +41 22 740 14 35			
For more detailed instructions, see the notes on th	e accompanying sheet.			
2. The applicant is hereby notified that no international search report will be established and that the declaration under Article 17(2)(a) to that effect and the written opinion of the International Searching Authority are transmitted herewith.				
3. With regard to the protest against payment of (an) ac	dditional fee(s) under Rule 40.2, the applicant is notified that:			
the protest together with the decision thereon has request to forward the texts of both the protest an	been transmitted to the International Bureau together with the applicant's d the decision thereon to the designated Offices.			
no decision has been made yet on the protest; the	applicant will be notified as soon as a decision is made.			
4. Reminders Shortly after the expiration of 18 months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the International Bureau as provided in Rules 90bis.1 and 90bis.3, respectively, before the completion of the technical preparations for international publication.				
International Bureau. The International Bureau will send a constraint of the preliminary examination report has been or is to be established before the expiration of 30 months from the priority date.	the applicant may submit comments on an informal basis on the written opinion of the International Searching Authority to the atternational Bureau. The International Bureau will send a copy of such comments to all designated Offices unless an international reliminary examination report has been or is to be established. These comments would also be made available to the public but not before the expiration of 30 months from the priority date.			
Within 19 months from the priority date, but only in respect of some designated Offices, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later); otherwise, the applicant must, within 20 months from the priority date, perform the prescribed acts for entry into the national phase before those designated Offices.  In respect of other designated Offices, the time limit of 30 months (or later) will apply even if no demand is filed within 19 months.				
	applicable time limits, Office by Office, see the PCT Applicant's Guide,			
Name and mailing address of the ISA/ US	Authorized officer Jameloh Dalin 1			
Mail Stop PCT, Attn: ISA/US Commissioner for Patents P.O. Box 1450	D. Margaret Seaman			
Alexandria, Virginia 22313-1450 Facsimile No. (703) 305-3230	Telephone No. 703-308-1235			
Form PCT/ISA/220 (January 2004)	(See notes on accompanying sheet)			
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# PATENT COOPERATION TREATY

# **PCT**

## INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

	Applicant's or agent's file reference 2307E-1439PC  FOR FURTHER see Form PCT/ISA/220 as well as, where applicable, item 5 below.					
	International application No. PCT/US05/07297  International filing date (day/month/year) 07 March 2005 (07.03.2005)  (Earliest) Priority Date (day/month/year) 05 March 2004 (05.03.2004)					
***	Applicant THE REGENTS OF HTE UNIVERSITY OF CALIFORNIA					
	This international search report has been prepared by this International Searching Authority and is transmitted to the applican according to Article 18. A copy is being transmitted to the International Bureau.  This international search report consists of a total of sheets.  It is also accompanied by a copy of each prior art document cited in this report.					
	<ol> <li>Basis of the Report         <ul> <li>With regard to the language, the international search was carried out on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.</li> </ul> </li> </ol>					
ļ	The international search was carried out on the basis of a translation of the international application furnished to this Authority (Rule 23.1(b)).					
	b. With regard to any nucleotide and/or amino acid sequence disclosed in the international application, see Box No. I.  Certain claims were found unsearchable (See Box No. II)					
	3. Unity of invention is lacking (See Box No. III)					
1	4. With regard to the title,					
)	the text is approved as submitted by the applicant.					
	the text has been established by this Authority to read as follows:					
}						
	5. With regard to the abstract,					
!	the text is approved as submitted by the applicant.					
	the text has been established, according to Rule 38.2(b), by this Authority as it appears in Box No. IV. The applicant					
	may, within one month from the date of mailing of this international search report, submit comments to this Authority.					
6	6. With regard to the drawings,					
	a. the figure of the drawings to be published with the abstract is Figure No  as suggested by the applicant.					
	as selected by this Authority, because the applicant failed to suggest a figure.					
	as selected by this Authority, because this figure better characterizes the invention.  b. In none of the figures is to be published with the abstract.					
L Fo	rm PCT/ISA/210 (first sheet) (January 2004)					

### INTERNATIONAL SEARCH REPORT

International application No.

PCT/US05/07297

A. CLASSIFICATION OF SUBJECT MATTER						
	IPC(7) : A61K 31/125; C07C 337/00; A61P 331/00					
US CL	: 514/582, 590, 238.7; 564/18, 20, 34; 544/162;					
	International Patent Classification (IPC) or to both na	itional classification and IPC				
	DS SEARCHED					
Minimum do	cumentation searched (classification system followed	by classification symbols)				
U.S. : 51	14/582, 590, 238.7; 564/18, 20, 34; 544/162; 546/153					
Documentation	on searched other than minimum documentation to the	outout that much do assessed as in to do do	4 - 6-14 1 - 4			
Documentati	on scarcined outer than infilling documentation to the	extent that such documents are included in	the neids searched			
Electronic da	ta base consulted during the international search (nam	e of data base and, where practicable, search	n terms used)			
West, East, S	TN: Registry, Chemical Abstracts	o or and once and, where practically, some	i torinis usou)			
C. DOCI	UMENTS CONSIDERED TO BE RELEVANT		<del></del>			
Category *	Citation of document, with indication, where a	and the second s	70.1			
P	LIS 6 907 240 A (COLEN et al.) 24 May 2005 (24 0)	appropriate, of the relevant passages	Relevant to claim No.			
P	US 6,897,240 A (COHEN et al) 24 May 2005 (24.0)	5.2005), abstract and formula I.	1-38			
P	US 6,903,237 A (YAMAGUCHI et al) 07 June 2005	(07.06.2005) shatmat	1 20			
1	1 03 0,703,237 X (174,0000111 et al) 07 Julie 2003	(07.00.2003), abstract.	1-38			
x	GREENBAUM D.C. Synthesis and Structure-Activi	ty Relationships of Parasiticidal	1-38			
	Thiosemicarbazone Cysteine Protease inhibitors aga	inst Plasmmodium falcinarum	1-36			
	Trypanosoma brucei and Trypanosoma cruzi, J. Med	I. Chem, May 2004, Vol 47 No.12, pages				
	3212-3219, page 3213.	, , , , , , , , , , , , , , , , , , , ,				
Ì						
X	DU X. Synthesis and Structure, J. Med. Chem., May	2002, Vol 45 No. 13, pages 2695-2707,	1-38			
	page 2696.					
i						
1						
	documents are listed in the continuation of Box C.	See patent family annex.				
* S <sub>I</sub>	pecial categories of cited documents:	"T" later document published after the inter				
"A" document	defining the general state of the art which is not considered to be of	date and not in conflict with the application principle or theory underlying the inver-				
particular	relevance	, , ,				
"E" earlier app	plication or patent published on or after the international filing date	"X" document of particular relevance; the cl considered novel or cannot be considered				
		when the document is taken alone	and an arrendite step			
	which may throw doubts on priority claim(s) or which is cited to he publication date of another citation or other special reason (as	"Y" document of particular relevance; the ci	simed invention cannot be			
specified)		considered to involve an inventive step	when the document is			
"O" document	referring to an oral disclosure, use, exhibition or other means	combined with one or more other such being obvious to a person skilled in the				
		•				
	published prior to the international filing date but later than the steel claimed	"&" document member of the same patent fa	amily			
		Data of mailing of the intermaliant				
Date of the actual completion of the international search  Date of mailing of the international search report			report			
14 June 2005	JØJUL 2004	İ				
Name and ma	iling address of the ISA/US	Authorized officer	Show hy			
	Stop PCT, Attn: ISA/US	D. Margaret Same-	gan in			
	missioner for Patents Box 1450	D. Margaret Seaman	77			
Alexandria, Virginia 22313-1450 Telephone No. 703-308-1235						
	Facsimile No. (703) 305-3230					

Form PCT/ISA/210 (second sheet) (January 2004)

### PATENT COOPERATION TREATY

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INTERNATIONAL SEARCH	HING AUTHORITY	hen		
To: ANNETTE S. PARENT TOWNSEND AND TOWN	SEND AND CREW LLP	# 3/)	PCT	
TWO EMBARCADERO CI SAN FRANCISCO, CA 94	ENTER, 8TH FLOOR		ITTEN OPINION OF THE DNAL SEARCHING AUTHORITY	
			(PCT Rule 43bis.1)	
62307E-12		Date of mailing (day/month/year) >	18 JUL 2003 .	
Applicant's or agent's file re	eference	FOR FURTHER	ACTION See paragraph 2 below	
International application No.	. International filing date	(day/month/year)	Priority date (day/month/year)	
PCT/US05/07297	07 March 2005 (07.03.20	005)		
	cation (IPC) or both national classification	ion and IPC	05 March 2004 (05.03.2004)	
IPC(7): A61K 31/125; C07C Applicant	2 337/00; A61P 331/00 and US Cl.: 514	/582, 590, 238.7; 564	1/18, 20, 34; 544/162; 546/153	
1	NIVERSITY OF CALIFORNIA			
1. This opinion contains in	dications relating to the following item	s:		
Box No. I	Basis of the opinion			
Box No. II	Priority			
Box No. III	Non-establishment of opinion with reg	gard to novelty, inven	tive step and industrial applicability	
Box No. IV	Lack of unity of invention	,		
Box No. V Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement				
Box No. VI				
Box No. VII	Box No. VII Certain defects in the international application			
Box No. VIII	Certain observations on the internation	al application		
2. FURTHER ACTION	4			
Authority other than this	y Examining Authority ("IPEA") exc	cept that this does to PEA has notified the	be considered to be a written opinion of the not apply where the applicant chooses an international Bureau under Rule 66.1 bis(b) ed.	
of Form PCT/ISA/220 or	ether, where appropriate, with amendn r before the expiration of 22 months fro	nents, before the exp	EA, the applicant is invited to submit to the iration of 3 months from the date of mailing thichever expires later.	
For further options, see I	Form PCT/ISA/220.	•		
3. For further details, see no	otes to Form PCT/ISA/220.			
Name and mailing address of		Authorized officer	<u> </u>	
Mail Stop PCT, Attn: Commissioner for Pate		D. Margaret Seama	an Janualey Shaken of	
P.O. Box 1450 Alexandria, Virginia 2			,	
Facsimile No. (703) 305-3230	)	Telephone No. 703	-308-1235	
Form PCT/ISA/237 (cover shee	et) (January 2004)		<del></del>	

resp to writte opinion DOCKETED BY

# · WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No. PCT/US05/07297

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Box No. V Reasoned statement under Rule 43 bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement					
1. Statement					
Novelty (N)	Claims NONE	YES			
	Claims 1-38	NO			
Inventive step (IS)	Claims NONE	YES			
	Claims 1-38	NO			
Industrial applicability (IA)	Claims 1-38	YES			
	Claims NONE	NONO			

#### 2. Citations and explanations:

Claims 1-38 lack novelty under PCT Article 33(2) as being anticipated by Du and Greenbaum. Both Du and Greenbaum teach compounds useful as antitrypanosomal therapy that anticipate the instant claims.

Claims 1-38 lack an inventive step under PCT Article 33(3) as being obvious over Du and Greenbaum. Both Du and Greenbaum teach generic compounds that are useful as antitrypanosomal therapy that render the intant claims obvious.

Claims 1-38 meet the criteria set out in PCT Article 33(4), and thus have industrial applicability because the subject matter claimed can be made or used in industry.

Form PCT/ISA/237 (Box No. V) (January 2004)

#### NOTESTOFORM PCT/ISA/220

These Notes are intended to give the basic instructions concerning the filing of amendments under Article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article," "Rule" and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions, respectively.

#### INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international publication. Furthermore, it should be emphasized that provisional protection is available in some States only.

#### What parts of the international application may be amended?

Under Article 19, only the claims may be amended

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Preliminary Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

When? Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

#### Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Scarching Authority (Rule 46.2).

Where a demand for international preliminary examination has been/is filed, see below.

How? Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions. Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

#### What documents must/may accompany the amendments?

#### Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.